# Low Adherence to KDIGO 2012 CKD Clinical Practice Guidelines Despite Clear Evidence of Utility

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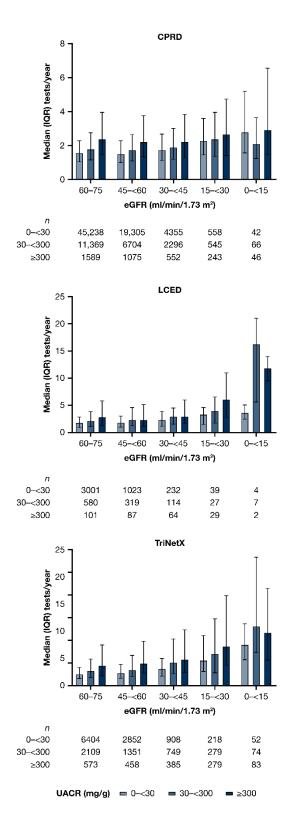
### **Supplementary Materials**

#### Supplementary Table 1. Ascertainment of clinical outcomes

Outcome	Method of ascertainment
≥50% eGFR decline	Two consecutive eGFR measures recorded ≥28 days
	apart (date of the second measurement is considered
	as the date of the adverse event)
Kidney failure	Progression to CKD stage 5 (sustained eGFR ≤15
	ml/min/1.73 m <sup>2</sup> ) or initiation of chronic RRT for >30
	days (2 dialysis codes 30 to 365 days apart) or kidney
	transplant
hHF	Hospitalization code for HF
Stroke	Diagnostic code
MI	Diagnostic code
All-cause mortality	ONS record or date of death of patient – derived using
	a CPRD algorithm (CPRD); death in hospital record
	(TriNetX)
Cardiovascular mortality	Primary diagnostic code on death record (CPRD only)

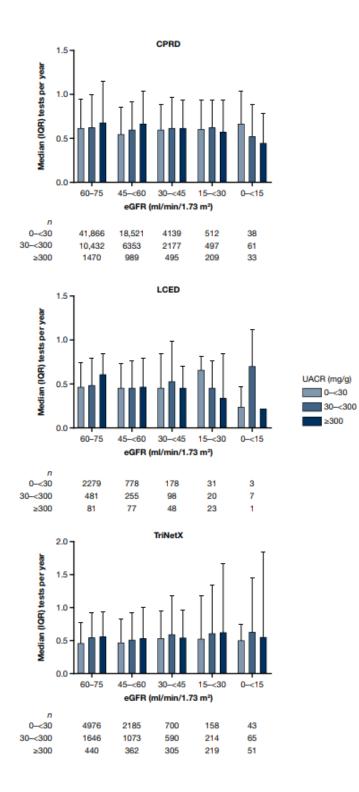
CKD, chronic kidney disease; CPRD, Clinical Practice Research Datalink; eGFR, estimated glomerular filtration rate; hHF, hospitalization for heart failure; MI, myocardial infarction; ONS, Office for National Statistics; RRT, renal replacement therapy.

#### Supplementary Figure 1. Frequency of eGFR testing per KDIGO category.



CPRD, Clinical Practice Research Datalink; eGFR, estimated glomerular filtration rate; IQR, interquartile range; KDIGO, Kidney Disease: Improving Global Outcomes; LCED, Limited Claims and Electronic Health Record Dataset; UACR, urinary albumin-to-creatinine ratio.

#### Supplementary Figure 2. Frequency of UACR testing per KDIGO category.



CPRD, Clinical Practice Research Datalink; eGFR, estimated glomerular filtration rate; IQR, interquartile range; KDIGO, Kidney Disease: Improving Global Outcomes; LCED, Limited Claims and Electronic Health Record Dataset; UACR, urinary albumin-to-creatinine ratio.

CPRD, LCED, and TriNetX data were from January 2008 to January 2020, January 2012 to June 2018, and January 2008 to March 2020, respectively.

#### Supplementary Figure 3. Patient comorbidities/complications by KDIGO

#### category at index.

			CPRD			LCED			TriNetX					CPRD			LCED			TriNetX			
AKI	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0– <30	30- <300	≥300	нк	UACR (mg/g)	0– <30	30- <300	≥300	0– <30	30- <300	≥300	0- <30	30- <300	≥300		
	60-75	0.9	1.7	3.8	2.5	3.6	10.3	2.5	5.4	9.6		60–75	0.3	0.5	1.3	1.7	2.5	1.7	1.3	1.3	4.4		
	45-<60	1.4	2.5	3.6	5.5	9.7	10.5	5.2	8.8	11.9		45-<60	0.3	0.6	1.0	2.6	5.0	5.3	2.0	2.0	3.6		
eGFR (ml/min/	30<45	2.9	4.4	6.8	9.7	17.5	31.0	11.9	18.3	20.1	eGFR (ml/min/	30-<45	0.7	1.0	1.0	4.7	7.1	5.6	3.1	3.1	6.9		
1.73 m²)	15-<30	7.9	12.1	15.3	29.5	21.4	38.7	28.8	29.7	28.2	1.73 m²)	15-<30	1.9	2.3	5.1	9.1	3.6	16.1	8.2	8.2	11.0		
	0-<15	46.5	32.4	27.1	50.0	87.5	66.7	28.8	30.3	33.7		0-<15	7.0	2.8	4.2	16.7	12.5	NA	19.2	19.2	9.0		
СНВ	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0 <30	30– <300	≥300	мі	UACR (mg/g)	0– <30	30- <300	≥300	0- <30	30– <300	≥300	0- <30	30- <300	≥300		
	60-75	16.7	19.0	20.8	19.0	29.1	34.5	15.9	21.9	22.1		60–75	7.0	7.7	9.6	4.8	8.9	5.2	3.9	6.7	6.5		
	45-<60	16.6	18.4	17.5	22.2	30.1	26.3	18.1	23.6	22.0	.055	45-<60	7.0	8.1	7.9	5.2	9.1	5.3	4.3	7.2	5.9		
eGFR (ml/min/ 1.73 m <sup>2</sup> )	30<45	19.7	20.9	19.8	27.2	31.7	25.4	22.5	29.3	30.6	eGFR (ml/min/ 1.73 m <sup>2</sup> )	30-<45	9.2	9.4	9.7	5.8	8.7	8.5	6.7	8.3	8.8		
1.101117	15-<30	23.5	17.2	18.4	31.8	14.3	22.6	29.2	29.1	28.2		15-<30	9.5	8.8	8.2	9.1	10.7	3.2	9.9	6.4	9.6		
	0-<15	20.9	19.7	12.5	16.7	50.0	33.3	15.4	18.4	23.6		0-<15	18.6	8.5	8.3	NA	37.5	NA	3.8	3.9	6.7		
T2D	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30– <300	≥300	Stroke	UACR (mg/g)	0– <30	30- <300	≥300	0- <30	30– <300	≥300	0- <30	30- <300	≥300		
	60-75	46.1	44.8	56.0	66.4	74.0	85.3	62.6	71.5	71.1		60–75	4.3	5.5	7.5	13.0	19.7	20.7	7.5	10.2	7.6		
.055	45-<60	22.1	25.3	38.7	65.7	73.2	83.2	61.1	71.7	73.3	eGFR (ml/min/ 1.73 m²)	45-<60	4.9	6.7	6.5	13.8	24.5	21.1	9.6	10.4	10.5		
eGFR (ml/min/ 1.73 m <sup>2</sup> )	30<45	18.3	20.5	26.3	69.6	75.4	76.1	62.8	66.9	73.5		30-<45	6.6	9.3	7.5	19.5	25.4	23.9	9.1	12.1	10.5		
1.101117	15-<30	22.3	18.4	24.7	75.0	67.9	64.5	60.1	59.8	66.8		15-<30	6.9	6.8	8.2	18.2	14.3	32.3	13.3	13.9	9.6		
	0-<15	23.3	21.1	27.1	66.7	50.0	33.3	32.7	25.0	48.3		0-<15	4.7	9.9	4.2	16.7	12.5	NA	7.7	5.3	7.9		
HF	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0 <30	30- <300	≥300	Cancer	UACR (mg/g)	0 <30	30- <300	≥300	0- <30	30– <300	≥300	0- <30	30- <300	≥300		
	60–75	3.5	5.5	6.6	5.3	11.7	13.8	5.6	9.1	11.7		60–75	16.2	17.2	16.1	4.5	6.6	1.7	3.8	3.9	3.6		
eGFR	45-<60	5.5	6.6	7.0	8.5	16.5	9.5	7.3	13.1	13.1	eGFR	45-<60	16.7	18.1	13.9	5.0	6.2	7.4	3.3	4.4	4.2		
(ml/min/ 1.73 m <sup>2</sup> )	30-<45	9.5	9.9	9.0	10.5	18.3	18.3	12.7	18.2	18.1	(ml/min/ 1.73 m <sup>2</sup> )	30-<45	18.5	21.2	15.4	7.4	10.3	12.7	4.7	3.4	3.2		
	15-<30	13.1	12.8	8.2	22.7	25.0	32.3	21.9	25.3	18.9		15-<30	18.4	21.5	20.4	13.6	10.7	<b>6</b> .5	4.7	4.7	2.7		
	0-<15	14.0	11.3	12.5	<mark>16.7</mark>	62.5	66.7	15.4	18.4	23.6		0-<15	18.6	16.9	14.6	16.7	12.5	NA	3.8	3.9	5.6		
нт	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300													
	60-75	57.6	63.4	68.9	82.4	88.9	89.7	74.0	79.9	78.1													
eGFR	45-<60	54.6	61.0	68.2	87.0	91.2	91.6	77.5	79.9	79.0													
(ml/min/ 1.73 m <sup>2</sup> )	30<45	62.2	62.7	65.0	93.4	92.9	94.4	77.0	74.1	78.2	2 Proportion of patients, %												
	15-<30	64.9	61.3	65.5	93.2	96.4	96.8	72.5	66.2	68.4			_	_									
	0-<15	76.7	67.6	56.2	100.0	75.0	66.7	40.4	44.7	66.3													

AKI, acute kidney injury; CHD, coronary heart disease; CPRD, Clinical Practice Research Datalink; eGFR, estimated glomerular filtration rate; HF, heart failure; HK, hyperkalemia; HT, hypertension; KDIGO, Kidney Disease: Improving Global Outcomes; LCED, Limited Claims and Electronic Health Record Dataset; MI, myocardial infarction; T2D, type 2 diabetes; UACR, urinary albumin-to-creatinine ratio. Values are % of patients; color coding is based on quartiles (Q) for each outcome within each database: green = Q1 and below; yellow = Q1 to Q2; orange = Q2 to Q3; red = Q3 and above.

#### Supplementary Figure 4. Prescription medications by KDIGO category at

#### index.

	114.00		CPRD			LCED			TriNetX					CPRD			LCED			TriNetX		
RAASia	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0– <30	30– <300	≥300	Insulin	UACR (mg/g)	0– <30	30- <300	≥300	0– <30	30- <300	≥300	0- <30	30- <300	≥300	
	60-75	50.7	57.9	67.8	62.1	74.0	78.4	44.3	53.8	58.4		60–75	4.4	7.5	17.2	11.6	21.1	29.3	16.8	30.3	38.0	
	45-<60	45.9	51.4	62.9	69.5	68.4	83.2	47.4	55.0	60.4		45-<60	2.3	4.2	11.3	11.8	20.9	27.4	18.5	29.6	38.8	
eGFR (ml/min/ 1.73 m <sup>2</sup> )	30<45	54.6	53.6	58.7	79.0	73.8	83.1	49.7	54.1	56.6	eGFR (ml/min/	30-<45	1.8	3.7	7.7	12.5	20.6	38.0	23.7	34.2	42.6	
1.73 m-)	15-<30	61.3	52.9	56.9	70.5	71.4	71.0	48.1	47.3	50.2	1.73 m²)	15-<30	4.0	3.5	4.7	27.3	14.3	48.4	29.2	31.1	45.5	
	0-<15	74.4	52.1	45.8	50.0	75.0	33.3	21.2	26.3	40.4		0-<15	9.3	1.4	4.2	16.7	37.5	66.7	26.9	15.8	37.1	
MRA	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0 <30	30– <300	≥300	MET	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30– <300	≥300	0- <30	30- <300	≥300	
	60–75	2.3	3.2	4.4	3.7	3.6	8.6	3.8	4.8	7.3		60–75	29.6	32.2	43.4	44.6	52.6	68.1	36.2	43.0	39.0	
	45-<60	3.5	3.9	3.6	4.5	8.0	7.4	4.6	5.8	6.1		45-<60	13.5	16.5	29.4	39.9	47.5	55.8	33.5	35.3	36.8	
eGFR (ml/min/ 1.73 m <sup>2</sup> )	30<45	5.1	5.5	3.9	6.6	4.8	4.2	7.9	8.0	7.6	eGFR (ml/min/ 1.73 m <sup>2</sup> )	30-<45	10.0	12.1	14.8	42.0	38.9	26.8	27.4	25.2	24.8	
1.75117	15-<30	8.8	7.2	5.5	9.1	NA	6.5	11.6	7.8	8.3	1.73111)	15-<30	11.7	10.3	12.2	34.1	32.1	19.4	21.9	15.2	16.9	
	0-<15	NA	8.5	6.2	16.7	NA	33.3	9.6	3.9	5.6		0-<15	14.0	15.5	14.6	33.3	12.5	33.3	13.5	7.9	9.0	
Diuretics	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30– <300	≥300	DPP4i	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30– <300	≥300	0- <30	30- <300	≥300	
	60–75	29.9	35.1	40.3	42.1	50.2	45.7	33.5	35.8	43.2		60–75	4.1	4.7	6.9	9.7	12.8	18.1	8.2	11.1	9.3	
.055	45-<60	35.7	39.6	41.0	53.0	54.0	51.6	38.5	42.0	47.7	eGFR (ml/min/ 1.73 m²)	45-<60	1.3	1.9	4.9	10.0	11.5	10.5	7.7	9.4	8.7	
eGFR (ml/min/ 1.73 m <sup>2</sup> )	30-<45	49.3	49.0	44.9	64.6	58.7	67.6	48.6	47.7	50.7		30-<45	1.0	1.4	3.2	14.0	14.3	14.1	9.7	10.7	8.3	
	15-<30	56.9	50.3	43.5	68.2	64.3	77.4	48.9	53.7	50.5		15-<30	1.0	1.4	3.1	9.1	7.1	9.7	7.7	10.1	8.0	
	0-<15	51.2	53.5	47.9	83.3	87.5	66.7	25.0	30.3	50.6		0-<15	4.7	4.2	6.2	16.7	25.0	NA	NA	6.6	5.6	
Statins	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	Steroids	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30– <300	≥300	0- <30	30- <300	≥300	
	60-75	56.6	58.5	62.8	63.9	70.4	74.1	45.9	50.7	50.5		60–75	8.9	10.1	10.4	15.8	14.6	16.4	0.1	<0.1	NA	
eGFR	45-<60	45.3	48.6	56.2	64.0	70.8	66.3	46.3	50.5	55.8	eGFR	45-<60	8.8	9.1	9.9	15.8	20.4	11.6	NA	NA	NA	
(ml/min/ 1.73 m <sup>2</sup> )	30-<45	44.4	43.9	44.7	67.7	65.9	67.6	49.3	52.6	55.6	(ml/min/ 1.73 m²)	30-<45	9.9	10.0	10.6	14.8	14.3	14.1	0.2	0.1	NA	
	15-<30	45.5	39.9	42.4	68.2	75.0	64.5	48.1	51.0	51.8		15-<30	11.2	10.2	9.8	11.4	17.9	25.8	NA	NA	NA	
	0-<15	51.2	45.1	41.7	33.3	62.5	100.0	21.2	31.6	48.3		0-<15	11.6	7.0	14.6	16.7	12.5	NA	NA	NA	NA	
ESA	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300												
	60-75	NA	NA	NA	0.1	0.2	NA	0.1	0.2	0.2												
eGFR	45-<60	<0.1	NA	NA	NA	0.6	NA	0.2	0.5	0.8	.5 Proportion of patients, %											
eGFR (ml/min/ 1.73 m <sup>2</sup> )	30-<45	<0.1	<0.1	0.3	NA	0.8	NA	0.5	1.1	0.5												
	15-<30	0.7	0.4	2.4	2.3	NA	6.5	0.9	2.4	5.6												
	0-<15	7.0	2.8	4.2	NA	25.0	NA	NA	6.6	10.1												

ACE, angiotensin-converting enzyme; ARB, angiotensin receptor blocker; CPRD, Clinical Practice Research Datalink; DPP4i, dipeptidyl peptidase-4; eGFR, estimated glomerular filtration rate; ESA, erythropoiesisstimulating agent; KDIGO, Kidney Disease: Improving Global Outcomes; LCED, Limited Claims and Electronic Health Record Dataset; MET, metformin; MRA, mineralocorticoid receptor antagonist; RAASi, renin-angiotensinaldosterone system inhibitor; UACR, urinary albumin-to-creatinine ratio.

<sup>a</sup>Includes ACE inhibitors and ARBs. Values are % of patients; color coding is based on quartiles (Q) for each parameter within each database: green = Q1 and below; yellow = Q1 to Q2; orange = Q2 to Q3; red = Q3 and above.

# Supplementary Figure 5. Laboratory parameters (in serum/whole blood) by KDIGO category at index.

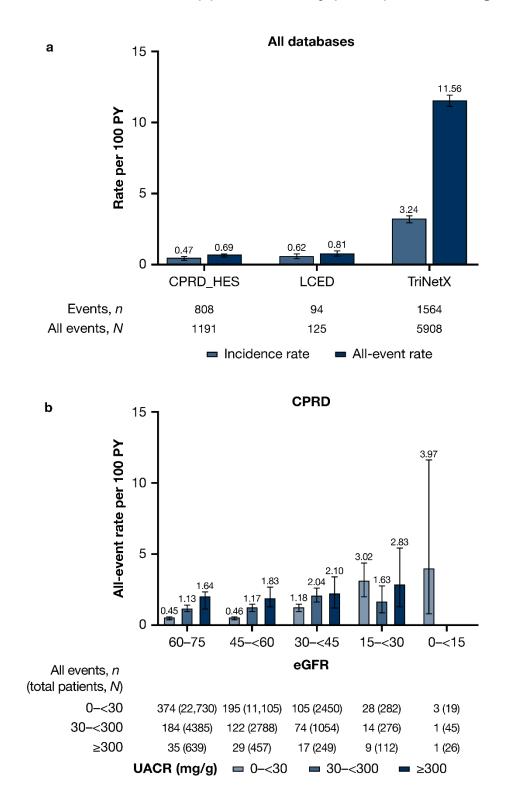
F	114.017		CPRD			LCED			TriNetX		Disastan i	114.00		CPRD		_	LCED			TriNetX	
Ferritin (ng/ml)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30– <300	≥300	Bicarbonate (mmol/L)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300
	60-75	125.9	125.1	139.5	142.0	173.7	128.3	150.6	217.6	194.1		60–75	26.6	26.5	26.2	26.4	26.3	26.0	26.6	26.4	26.2
	45<60	129.1	140.1	154.1	146.2	249.6	102.4	<b>1</b> 47.6	191.4	216.7		45-<60	26.4	26.6	26.2	26.3	26.1	25.2	26.3	26.1	25.5
eGFR (ml/min/	30<45	151.4	163.1	186.0	174.8	93.5	130.4	178.0	202.2	210.6	eGFR (ml/min/	30-<45	26.2	25.4	25.2	25.7	25.0	24.4	25.9	25.4	25.4
1.73 m²)	15-<30	166.2	180.4	220.7	124.9	52.7	215.8	128.4	237.8	246.3	1.73 m²)	15-<30	25.2	23.9	23.7	24.0	25.3	24.0	24.9	24.8	23.7
	0-<15	181.4	146.5	321.6	13.3	187.9	127.5	286.3	301.1	184.4		0-<15	22.3	20.9	22.7	20.2	21.5	24.7	24.0	23.1	22.0
Albumin (g/dL)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	Uric acid (µmol/L)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300
	60-75	4.3	4.2	4.0	4.3	4.3	4.1	4.1	4.1	3.8		60–75	370.1	376.5	404.5	359.1	368.2	462.5	351.8	355.8	347.7
	45-<60	4.2	4.2	4.0	4.3	4.2	3.9	4.1	4.0	3.8		45-<60	388.8	395.2	419.6	368.4	390.6	392.9	368.9	385.9	393.5
eGFR (ml/min/ 1.73 m²)	30<45	4.2	4.1	3.9	4.2	4.2	4.0	4.0	3.9	3.7	eGFR (ml/min/ 1.73 m²)	30-<45	423.4	424.3	425.8	366.4	377.6	424.3	395.6	386.9	409.9
	15-<30	4.1	4.0	3.8	4.1	3.9	3.9	3.9	3.9	3.7		15-<30	424.6	424.8	456.8	444.8	452.1	455.0	399.1	404.8	420.1
	0-<15	3.8	3.9	3.7	4.0	3.9	NA	3.8	3.7	3.7		0-<15	398.5	418.8	NA	NA	466.9	398.5	436.2	425.3	439.0
K⁺ (mmol/L)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0 <30	30– <300	≥300	Hb (g/dL)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300
	6075	4.5	4.5	4.5	4.3	4.3	4.4	4.2	4.2	4.3		60–75	13.9	13.7	13.4	13.8	13.7	13.6	13.6	13.4	13.2
eGFR	45-<60	4.5	4.5	4.6	4.3	4.3	4.3	4.2	4.3	4.3	eGFR	45-<60	13.6	13.4	13.0	13.5	13.5	13.1	13.2	12.9	12.7
(ml/min/ 1.73 m²)	30<45	4.6	4.6	4.6	4.5	4.5	4.4	4.3	4.3	4.4	(ml/min/ 1.73 m²)	30-<45	12.9	12.7	12.7	13.0	12.8	12.7	12.6	12.3	12.2
	15-<30	4.7	4.7	4.8	4.4	4.6	4.6	4.5	4.5	4.5		15-<30	12.2	11.9	11.9	11.9	12.0	11.3	11.9	11.7	11.5
	0-<15	4.9	4.8	4.9	4.8	4.6	4.2	4.3	4.3	4.5		0-<15	11.2	10.9	11.2	11.2	9.7	11.7	11.1	10.9	10.7
Creatinine (mg/dL)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30– <300	≥300	Calcium (mg/dL)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300
	60–75	1.0	1.0	1.1	1.0	1.0	1.1	1.0	1.0	1.1		60–75	9.4	9.4	9.3	9.5	9.5	9.5	9.4	9.4	9.4
eGFR	45-<60	1.2	1.2	1.3	1.2	1.4	1.3	1.2	1.2	1.3	eGFR	45-<60	9.5	9.4	9.3	9.5	9.5	9.5	9.5	9.5	9.3
(ml/min/ 1.73 m²)	30<45	1.4	1.5	1.6	1.5	1.5	2.2	1.5	1.6	1.6	(ml/min/ 1.73 m <sup>2</sup> )	30-<45	9.4	9.4	9.3	9.5	9.5	9.5	9.4	9.4	9.2
	15-<30	2.1	2.2	2.5	2.1	2.2	2.6	2.2	2.3	2.4		15-<30	9.5	9.2	9.2	9.5	9.3	9.2	9.4	9.3	9.1
	0-<15	4.7	4.3	4.8	3.8	4.0	4.1	5.6	5.7	5.1		0-<15	8.9	9.1	9.1	9.0	8.6	9.1	8.8	8.9	8.9
BNP (pg/ml)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	Sodium (mmol/L)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300
	60-75	NA	NA	NA	170.3	414.1	143.0	224.7	215.7	275.7		60–75	140. <b>1</b>	139.7	139.4	139.7	139.4	139.5	139.3	138.8	138.4
eGFR	45<60	NA	NA	NA	245.3	244.9	63.5	218.4	309.3	302.1	eGFR	45-<60	140.1	139.9	139.4	139.8	139.2	139.4	139.2	138.8	138.6
(ml/min/ 1.73 m²)	30<45	NĂ	NA	NA	102.6	511.2	78.7	342.6	345.2	493.8	(ml/min/ 1.73 m²)	30-<45	139.7	139.6	139.3	139.4	139.4	138.6	139.0	138.7	138.9
	15-<30	NA	NA	NA	452.0	801.3	112.7	371.4	447.9	522.2		15-<30	139.4	139.4	139.5	138.7	140.0	138.8	137.7	138.5	138.1
	0-<15	NA	NA	NA	NA	291.5	358.0	204.4	593.9	916.0		0-<15	138.6	138.6	138.3	136.2	139.0	142.0	137.1	137.4	138.0

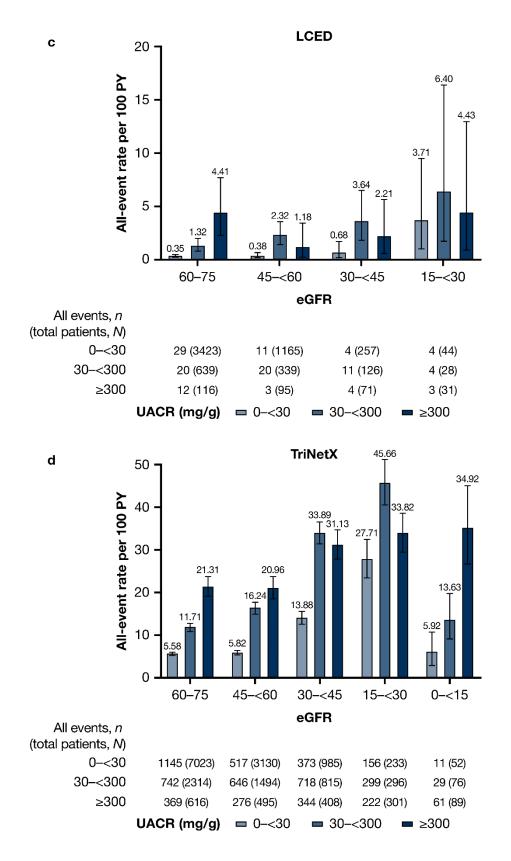
CRP (mg/L)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	Phosphate (mmol/L)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	
	60-75	10.4	15.0	17.9	13.4	18.8	24.5	8.3	7.4	9.9		60-75	1.1	1.1	1.1	1.1	1.0	1.1	NA	NA	NA	
	45-<60	12.3	17.5	18.7	15.5	26.7	35.4	9.4	16.0	13.2		45-<60	1.1	1.1	1.1	1.1	1.1	1.2	NA	NA	NA	
eGFR (ml/min/ 1.73 m²)	30<45	21.3	20.9	26.1	7.1	4.0	12.6	11.7	12.7	5.6	eGFR (ml/min/ 1.73 m²)	30<45	1.1	1.1	1.2	1.0	1.1	1.2	NA	NA	NA	
1.75111	15-<30	27.6	31.8	31.3	NA	51.9	3.0	18.9	7.6	8.9	1.75117	15-<30	1.2	1.2	1.2	1.0	1.1	1.3	NA	NA	NA	
	0-<15	35.4	33.4	83.8	NA	NA	NA	88.7	11.5	5.0		0-<15	1.5	1.4	1.6	2.2	1.5	1.2	NA	NA	NA	
LDL (mmol/L)	UACR (mg/g)	0– <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	WBC Count (10*9/L)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0 <30	30- <300	≥300	
	60–75	2.7	2.6	2.6	2.5	2.4	2.5	2.4	2.3	2.5		60–75	7.4	7.6	8.1	8.4	8.7	8.0	NA	NA	NA	
	45-<60	2.9	2.7	2.7	2.5	2.4	2.5	2.4	2.3	2.4		45-<60	7.3	7.7	8.1	7.9	10.3	7.8	NA	NA	NA	
eGFR (ml/min/	30<45	2.8	2.7	2.8	2.4	2.4	2.7	2.4	2.2	2.4	eGFR (ml/min/	30-<45	7.6	7.9	8.1	7.5	7.4	NA	NA	NA	NA	
1.73 m²)	15-<30	2.8	2.7	2.9	2.5	2.1	2.8	2.4	2.2	2.3	1.73 m²)	15-<30	7.9	8.3	8.3	NA	NA	6.6	NA	NA	NA	
	0-<15	2.2	3.1	3.1	2.1	2.7	2.1	2.3	2.3	2.4		0-<15	9.2	9.3	8.8	NA	17.2	NA	NA	NA	NA	
HDL (mmol/L)	UACR (mg/g)	0– <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	lron (µg/dL)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	
	60-75	1.3	1.3	1.3	1.3	1.2	1.1	1.3	1.2	1.2		60-75	84.4	77.9	72.7	81.4	69.3	75.3	70.8	68.3	70.9	
eGFR	45-<60	1.4	1.4	1.3	1.3	1.2	1.2	1.3	1.2	1.2	eGFR	45-<60	82.0	71.8	59.5	80.6	70.6	73.6	66.2	63.2	60.2	
(ml/min/ 1.73 m²)	30-<45	1.4	1.4	1.3	1.3	1.2	1.2	1.3	1.2	1.2	eGFR (ml/min/ 1.73 m²)	30-<45	72.5	72.9	62.5	73.2	73.8	63.7	70.7	66.6	65.1	
, ,	15-<30	1.4	1.3	1.2	1.2	1.1	1.3	1.3	1.2	1.1		15-<30	74.6	59.3	69.2	109.3	55.0	63.1	64.1	59.5	57.0	
	0-<15	1.2	1.2	1.4	1.1	0.8	1.4	1.1	1.0	1.3		0-<15	NA	73.5	49.7	73.0	40.8	48.0	74.4	50.9	55.6	
Transferrin saturation (%)	UACR (mg/g)	0– ⊲30	30- ⊲300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	Parathyrin parathyroid (pg/ml)	UACR (mg/g)	0- <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300	
	60–75	27.3	26.8	27.1	24.7	25.0	18.3	22.5	23.1	23.9		60–75	76.0	63.8	55.6	NA	NA	NA	NA	NA	NA	
eGFR	45-<60	28.0	24.2	24.8	24.6	22.0	21.3	22.1	19.9	21.1	eGFR	45-<60	81.7	80.4	161.3	NA	NA	NA	NA	NA	NA	
eGFR (ml/min/ 1.73 m²)	30<45	23.5	28.1	21.2	23.5	24.7	17.9	22.3	22.8	23.1	(ml/min/ 1.73 m²)	30-<45	77.9	84.4	110.3	NA	NA	NA	NA	NA	NA	
	15-<30	33.4	23.8	40.3	52.9	24.2	22.5	20.9	21.8	20.3		15-<30	88.4	272.3	129.3	NA	NA	NA	NA	NA	NA	
	0-<15	NA	NA	NA	22.3	18.5	18.3	24.5	20.0	25.5		0-<15	147.1	194.7	194.2	NA	NA	NA	NA	NA	NA	
TIBC (µg/dL)	UACR (mg/g)	0– <30	30- <300	≥300	0- <30	30- <300	≥300	0- <30	30- <300	≥300												
	60–75	314.3	318.1	301.6	337.9	319.9	378.4	340.9	329.1	318.8												
eGFR	45-<60	302.7	316.1	292.3	332.6	329.6	329.0	328.5	326.1	303.6												
eGFR (ml/min/ 1.73 m²)	30<45	281.5	306.9	277.5	307.8	330.2	326.6	326.6	309.5	293.3	incur value											
	15-<30	247.2	252.3	221.4	266.0	257.5	284.1	311.7	281.2	270.7												
	0<15	NA	NA	NA	NA	245.3	277.5	267.8	262.8	245.6	56											

BNP, brain natriuretic peptide; CPRD, Clinical Practice Research Datalink; CRP, C-reactive protein; eGFR, estimated glomerular filtration rate; Hb, hemoglobin; HDL, high density lipoprotein cholesterol; K<sup>+</sup>, potassium; KDIGO, Kidney Disease: Improving Global Outcomes; LCED, Limited Claims and Electronic Health Record Dataset; LDL, low density lipoprotein cholesterol; TIBC, total iron binding capacity; UACR, urinary albumin-to-creatinine ratio; WBC, white blood cell.

Mean values are shown; color coding is based on quartiles (Q) for each parameter within each database: green = Q1 and below; yellow = Q1 to Q2; orange = Q2 to Q3; red = Q3 and above.

Supplementary Figure 6. Incidence rates<sup>a</sup> and all-event rates<sup>b</sup> per 100 PY of hospitalization for heart failure: (a) overall and by (b, c, d) KDIGO category.





CI, confidence interval; CPRD, Clinical Practice Research Datalink; eGFR, estimated glomerular filtration rate; HF, heart failure; HES, hospital episode statistics; LCED, Limited Claims and Electronic Health Record Dataset; PY, person-years; UACR, urinary albumin-to-creatinine ratio. Error bars represent 95% CI. Note: TriNetX does not record a primary diagnosis during hospitalization, meaning that data show inpatient events where HF is recorded, and therefore, not necessarily the primary diagnosis. <sup>a</sup>Calculated as the number of specific events that occur during patient follow-up time at risk (time in the study until first event or loss to follow-up).

<sup>b</sup>Calculated as the total number of events that occurred during patient follow-up time at risk (patients' time until loss to follow-up).

#### Supplementary Figure 7. Incidence rates per 100 PY of clinical outcomes<sup>a</sup>

#### during follow-up by KDIGO category (sensitivity analysis).

≥50%	UACR		CPRD			LCED			TriNetX	
eGFR decline	(mg/g)	0-<30	30-<300	≥300	0-<30	30-<300	≥300	0-<30	30-<300	≥300
	60–75	0.12 (0.0 <del>9-</del> 0.15)	0.44 (0.33–0.58)	2.62 (1.84–3.61)	0.08 (0.02–0.18)	0.47 (0.15–1.11)	1.67 (0.34–4.88)	0.19 (0.13–0.27)	0.59 (0.3 <del>9-</del> 0.85)	2.88 (2.02–3.99)
	45-<60	0.13 (0.09–0.18)	0.39 (0.27–0.56)	1.76 (1.06–2.74)	0.18 (0.05–0.46)	NA	3.32 (1.22–7.23)	0.31 (0.19–0.47)	0.73 (0.45–1.12)	4.19 (2.98–5.73)
eGFR (ml/min/ 1.73 m²)	30-<45	0.15 (0.07–0.27)	0.46 (0.24–0.80)	1.63 (0.78–3.00)	NA	NA	4.83 (1.77–10.52)	0.59 (0.30–1.03)	1.36 (0.84–2.07)	4.76 (3.33–6.59)
	15-<30	0.88 (0.32–1.92)	NA	3.11 (1.34–6.13)	NA	NA	NA	1.68 (0.67–3.46)	NA	8.03 (5.66–11.08)
	0-<15	NA	NA	NA	NA	NA	NA	NA	1.71 (0.35–4.99)	4.93 (1.98–10.16)
Kidney failure	UACR (mg/g)	0-<30	30-<300	≥300	0-<30	30-<300	≥300	0-<30	30-<300	≥300
	60–75	0.01 (0.00–0.02)	0.03 (0.01–0.09)	1.17 (0.68–1.88)	0.05 (0.01–0.13)	NA	NA	0.04 (0.01–0.08)	0.25 (0.13–0.44)	1.01 (0.54–1.73)
	45-<60	0.02 (0.01–0.04)	0.12 (0.05–0.22)	1.37 (0.77–2.26)	0.13 (0.03–0.39)	NA	2.70 (0.88–6.29)	0.04 (0.01–0.13)	0.28 (0.12–0.55)	2.20 (1.36–3.37)
eGFR (ml/min/ 1.73 m²)	30-<45	0.07 (0.02–0.17)	0.27 (0.11–0.55)	3.45 (2.14–5.27)	NA	NA	4.73 (1.74–10.29)	0.34 (0.14–0.71)	1.23 (0.74–1.91)	4.92 (3.47–6.79)
	15-<30	1.80 (0.93–3.15)	2.47 (1.41–4.02)	11.88 (7.83–17.28)	NA	6.40 (1.32–18.71)	22.85 (11.40–40.88)	3.67 (2.06–6.06)	2.11 (1.01–3.88)	22.13 (17.77–27.23)
	0–<15	NA	14.33 (7.15–25.64)	4 <b>1</b> .71 (23.34–68.79)	NA	NA	NA	1.94 (0.40–5.66)	7.97 (4.24–13.62)	48.33 (35.51–64.26)
hHF	UACR (mg/g)	0-<30	30-<300	≥300	0-<30	30-<300	≥300	0-<30	30-<300	≥300
	60–75	0.28 (0.24–0.33)	0.90 (0.73–1.09)	1.33 (0.80–2.07)	0.28 (0.16–0.44)	1.05 (0.53–1.88)	3.98 (1.60–8.20)	1.67 (1.47–1.88)	3.94 (3.38–4.56)	5.44 (4.20–6.94)
	45-<60	0.45 (0.38–0.53)	0.82 (0.63–1.05)	1.58 (0.92–2.53)	0.36 (0.15–0.7)	1.68 (0.80–3.09)	NA	2.12 (1.78–2.50)	4.51 (3.74–5.38)	6.63 (5.05–8.56)
eGFR (ml/min/ 1.73 m²)	30-<45	0.92 (0.71–1.18)	1.80 (1.33–2.40)	1.66 (0.79–3.04)	NA	2.95 (1.08–6.42)	2.41 (0.50–7.05)	3.83 (3.00–4.80)	8.32 (6.88–9.96)	13.22 (10.60–16.28)
	15-<30	2.85 (1.72–4.45)	1.68 (0.84–3.01)	1.92 (0.62–4.48)	3.46 (0.71–10.12)	NA	6.25 (1.29–18.27)	9.29 (6.51–12.86)	15.37 (11.78–19.70)	13.94 (10.67–17.91)
	0-<15	NA	NA	NA	NA	NA	NA	3.31 (1.07–7.72)	3.50 (1.28–7.61)	12.75 (7.43–20.41)
Stroke	UACR (mg/g)	0-<30	30-<300	≥300	0-<30	30-<300	≥300	0-<30	30-<300	≥300
	60–75	0.78 (0.71–0.86)	1.09 (0.91–1.30)	2.34 (1.61–3.29)	5.45 (4.88–6.08)	10.26 (8.28–12.57)	10.66 (6.21–17.06)	4.12 (3.79–4.46)	5.59 (4.91–6.34)	5.40 (4.17–6.89)
	45-<60	0.97 (0.86–1.09)	1.34 (1.09–1.63)	1.66 (0.98–2.63)	6.83 (5.73–8.07)	12.16 (9.33–15.59)	11.01 (6.41–17.62)	5.34 (4.78–5.94)	6.64 (5.68–7.70)	8.18 (6.38–10.34)
eGFR (ml/min/ 1.73 m²)	30-<45	1.61 (1.32–1.94)	1.94 (1.44–2.56)	1.99 (1.03–3.47)	10.13 (7.20–13.85)	14.56 (9.42–21.49)	11.20 (5.79–19.56)	6.29 (5.19–7.54)	9.28 (7.74–11.05)	8.33 (6.33–10.77)
	15-<30	2.99 (1.83–4.62)	1.99 (1.06–3.41)	2.69 (1.08–5.53)	9.27 (3.73–19.11)	11.83 (3.84–27.61)	13.23 (4.86–28.80)	8.15 (5.50–11.63)	10.68 (7.76–14.33)	6.86 (4.66–9.73)
	0-<15	NA	NA	NA	NA	NA	NA	1.97 (0.41–5.76)	6.84 (3.41–12.23)	6.99 (3.35–12.85)

мі	UACR (mg/g)	0<30	30-<300	≥300	0-<30	30-<300	≥300	0-<30	30-<300	≥300
	60–75	1.36 (1.27–1.46)	2.10 (1.84–2.38)	2.64 (1.86–3.64)	2.33 (1.96–2.73)	5.06 (3.76–6.68)	9.55 (5.46–15.51)	2.42 (2.18–2.68)	4.20 (3.63–4.85)	4.70 (3.56–6.08)
	45-<60	1.38 (1.25–1.52)	1.87 (1.57–2.20)	3.42 (2.40–4.74)	3.29 (2.57–4.16)	5.87 (4.04–8.24)	3.38 (1.24–7.35)	2.83 (2.44–3.27)	4.44 (3.68–5.31)	6.41 (4.84–8.33)
eGFR (ml/min/ 1.73 m²)	30-<45	2.00 (1.67–2.37)	1.98 (1.47–2.60)	2.68 (1.53–4.36)	5.94 (3.84–8.77)	6.92 (3.78–11.61)	6.84 (2.95–13.48)	3.74 (2.92–4.70)	6.98 (5.68–8.50)	8.58 (6.56–11.02)
	15-<30	3.32 (2.08–5.03)	3.32 (2.06–5.08)	2.34 (0.86–5.09)	5.90 (1.92–13.77)	NA	6.13 (1.26–17.92)	5.88 (3.72–8.82)	6.56 (4.40–9.43)	8.71 (6.20–11.91)
	0-<15	NA	3.34 (0.69–9.77)	NA	NA	NA	NA	NA	2.95 (0.96–6.88)	7.06 (3.39–12.99)
All-cause mortality	UACR (mg/g)	0<30	30<300	≥300	0<30	30-<300	≥300	0-<30	30-<300	≥300
	60–75	1.22 (1.13–1.31)	2.62 (2.33–2.93)	4.39 (3.38–5.61)	NA	NA	NA	0.84 (0.70–0.99)	1.50 (1.17–1.89)	1.95 (1.26–2.88)
	45-<60	2.09 (1.93–2.26)	4.17 (3.73–4.66)	5.17 (3.92–6.70)	NA	NA	NA	0.94 (0.72–1.20)	2.48 (1.93–3.13)	2.31 (1.45–3.50)
eGFR (ml/min/ 1.73 m²)	30-<45	4.46 (3.97–4.99)	7.74 (6.72 <b>–8.88</b> )	6.31 (4.48–8.62)	NA	NA	NA	1.03 (0.64–1.58)	2.40 (1.69–3.31)	2.59 (1.58–4.00)
	15-<30	9.91 (7.69 <b>–12</b> .56)	12.54 (9.98–15.54)	10.61 (7.05–15.34)	NA	NA	NA	3.84 (2.19–6.23)	3.62 (2.11–5.80)	3.27 (1.87–5.31)
	0-<15	5.85 (1.21–17.10)	14.16 (7.54–24.22)	NA	NA	NA	NA	1.97 (0.41–5.76)	2.27 (0.62–5.82)	5.33 (2.30–10.51)
CV mortality	UACR (mg/g)	0-<30	30-<300	≥300	0-<30	30-<300	≥300	0-<30	30-<300	≥300
	60–75	0.29 (0.25–0.33)	0.71 (0.56–0.88)	1.03 (0.58–1.70)	NA	NA	NA	NA	NA	NA
	45-<60	0.43 (0.36–0.51)	0.82 (0.63–1.05)	1.81 (1.11–2.80)	NA	NA	NA	NA	NA	NA
eGFR (ml/min/ 1.73 m²)	30-<45	1.18 (0.94–1.47)	1.59 (1.15–2.15)	1.29 (0.56–2.55)	NA	NA	NA	NA	NA	NA
	15-<30	2.04 (1.11–3.42)	1.96 (1.05–3.36)	1.89 (0.62–4.42)	NA	NA	NA	NA	NA	NA
	0-<15	NA	3.27 (0.67–9.55)	NA	NA	NA	NA	NA	NA	NA

CI, confidence interval; CPRD, Clinical Practice Research Datalink; CV, cardiovascular; eGFR, estimated glomerular filtration rate; hHF, hospitalization for heart failure; KDIGO, Kidney Disease: Improving Global Outcomes; LCED, Limited Claims and Electronic Health Record Dataset; MI, myocardial infarction; NA, not available (owing to low patient/event numbers); PY, patient-years; UACR, urinary albumin-to-creatinine ratio. Data are incidence rate (95% CI), calculated as the number of specific events that occur during patient follow-up time at risk (time in the study until first event or loss to follow-up). Color coding is based on odds ratio quartile (Q) for each outcome within each database: green = Q1 and below; yellow = Q1 to Q2; orange = Q2 to Q3; red = Q3 and above.

<sup>a</sup>Mortality data were not available for US LCED and incidence rates for some outcomes were not available where there were low patient or event numbers (e.g., LCED eGRF <15 ml/min/1.73 m<sup>2</sup>).

# How Clinical Guidelines for **CHRONIC KIDNEY DISEASE** Have Been Used in the Real World Since 2012

### WHAT WAS THE FOCUS **OF THE STUDY?**

WHAT WAS THE AIN

Estimated glomerular filtration rate (eGFR) and urinary albumin-to-creatinine ratio (UACR) are two key tests for chronic kidney disease (CKD) that can help doctors, patients, and caregivers predict and manage the risk of CKD complications.

#### eGFR test

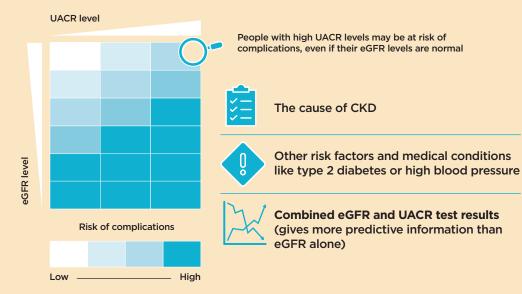
Shows how well a person's kidneys are functioning by measuring a waste product called creatinine in the blood.

#### **UACR** test

.....

Measures whether a type of protein called albumin has leaked from the kidnevs into the urine due to kidney damage.

#### We can predict the risk of serious complications of CKD by looking at:



RESEARCHERS WANTED TO

treatments, and

large databases of

the anonymized

medical records of

patient medical records:

**Researchers** looked at

To understand more about the

characteristics, medication

people living with CKD.

use, and health outcomes of

How do characteristics. outcomes change for patients with less or more severe CKD as defined in the guidelines?

How often are eGFR and UACR tests carried out on people living with CKD?

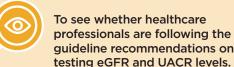
**OF THIS STUDY** 

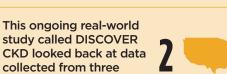
FIND OUT

Patients with type 1 diabetes, or who had received a kidney transplant or were receiving dialysis were not included in this study.

#### Patients eligible for the study







?

from the US and from the UK



adults

with CKD

#### The following health outcomes were tested:

# 50%

Kidney disease getting worse (a sustained 50% reduction in eGFR or development of kidney failure).

Across-database range

Of

Heart problems (heart attack, stroke, or hospitalization for heart failure).



Loss of life from any cause, or loss of life caused by disease of the heart and blood vessels (cardiovascular disease).

Years

had 1 or more UACR

measure(s) and

in the study

could be included

## WHAT WERE THE RESULTS OF THE STUDY?

Many people with CKD included in the study were categorized by the clinical guidelines as being in the low or moderately increased risk groups for CKD complications. The proportion of participants in each risk group was:

#### **CKD** risk group Across-database range 38.3-53.9% 28.4-32.5% 11.2-17.0% 5.9-15.1% Low risk Moderately **High risk** Very high risk increased risk In this study... PARTICIPANTS MOST PARTICIPANTS THE AVERAGE TIME THAT THAT WERE MEN WERE AGED BETWEEN **PATIENTS WERE** FOLLOWED IN THE DATABASE FOR WAS

Years

people with 2

in the medical

eGFR measures

databases, only

#### As eGFR decreased and UACR increased:



There was increased risk of kidney disease getting worse, heart problems, and loss of life. Prescriptions for some CKD medicines increased.



Occurrence of other medical conditions, such as type 2 diabetes and high blood pressure, tended to increase.

# eGFR and UACR were tested less often than the recommended once per year for low risk patients, and 3 to 4 or more times per year for very high risk patients

The average number of tests per year were:



### WHAT DO THESE RESULTS MEAN FOR PEOPLE LIVING WITH CKD?



eGFR and UACR levels are linked with the expected health outcomes of people living with CKD.



More regular testing for all people living with CKD is recommended for monitoring the risk of kidney disease getting worse, heart disease, and loss of life.



You can read the full medical paper by James G et al. Low Adherence to KDIGO 2012 CKD Clinical Practice Guidelines Despite Clear Evidence of Utility.



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